

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Elizabeth M. Denholm, Yong-Qing Lin and Paul J. Silver

Serial No.: Continuation of 09/715,965

Art Unit: Not Yet Assigned

Filed: July 18, 2003

Examiner: Not Yet Assigned

For: *ATTENUATION OF TUMOR GROWTH, METASTASIS AND ANGIOGENESIS*

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

**INFORMATION DISCLOSURE STATEMENT**

Sir:

Pursuant to 37 C.F.R. §1.56 and 37 C.F.R. §1.97, Applicants submit an Information Disclosure Statement, including five (5) pages of Form PTO-1449. All of the documents cited below were cited by or submitted to the Patent Office in Application Serial No. 09/715,965, filed November 17, 2000, to which the present application claims priority. Pursuant to 37 C.F.R. §1.98(d), Applicants are not enclosing copies of these publications. Copies will be provided upon request, however.

This Information Disclosure Statement is being filed under 37 C.F.R. § 1.97(b) prior to a first Office Action on the merits. It is believed that no fee is required with this submission. However, should a fee be required, the Commissioner is hereby authorized to charge any required fees to Deposit Account No. 50-1868.

### U.S. Patents

<u>Number</u>	<u>Issue Date</u>	<u>Patentee</u>	<u>Class/Subclass</u>
4,696,816	09-29-1987	Brown	424/94
5,567,417	10-22-1996	Sasisekharan et al.	424/94.5
5,945,403	08-31-1999	Folkman et al.	514/21

### Foreign Documents

<u>Number</u>	<u>Publication Date</u>	<u>Patentee</u>	<u>Country</u>
51075042	06-29-1979	Ono Pharm Co. Ltd.	JP
WO 96/01894	01-25-1996	IBEX Technologies	PCT
WO 96/01648	01-25-1996	IBEX Technologies	PCT
WO 96/08559	03-21-1996	Cardiac CRC Nominees PTY, LTD.	

### Publications

CRUM, et al., "A new class of steroids inhibits angiogenesis in the presence of heparin or a heparin fragment," *Science* 230(4732): 1375-1378 (1985).

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FOLKMAN, "Successful treatment of an angiogenic disease," *N. Engl. J. Med.* 320(18): 1211-1212 (1989).

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FOLKMAN, et al., "Control of angiogenesis with synthetic heparin substitutes," *Science* 243(4897): 1490-1493 (1989).

FORRESTER, et al., "A paradigm for restenosis based on cell biology: clues for the development of new preventive therapies," *J. Am. Coll. Cardiol.* 17(3):758-769 (1991).

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HENKE, et. al., "CD44-related chondroitin sulfate proteoglycan, a cell surface receptor implicated with tumor cell invasion, mediates endothelial cell migration on fibrinogen and invasion into a fibrin matrix," *J. Clin. Invest.* 97(11):2541-2552 (1996).

INGBER, et al., "*Inhibition of angiogenesis through modulation of collagen metabolism*," *J. Lab. Invest.* 59: 44-51 (1989).

INGBER, et al., "A possible mechanism for inhibition of angiogenesis by angiostatic steroids: induction of capillary basement membrane dissolution," *Endocrinol.* 119(4): 1768-1775 (1986).

INGBER, et al., "Mechanochemical switching between growth and differentiation during fibroblast growth factor-stimulated angiogenesis in vitro: role of extracellular matrix," *J. Cell. Biol.* 109(1): 317-330 (1989).

JACKSON et. al., "Glycosaminoglycans: molecular properties, protein interactions, and role in physiological processes," *Physiol. Rev.* 71(2):481-530 (1991).

JIN-INCHI, et al., "Inhibition of experimental metastasis of murine Lewis lung carcinoma by an inhibitor of glucosylceramide synthase and its possible mechanism of action," *Cancer Res.* 50:6731-6737 (1990).

LIDA, et al., "Cell surface chondroitin sulfate proteoglycans in tumor cell adhesion, motility and invasion," *Sem. Cancer Biol.* 7:155-162, (1996).

LINHARDT, et al., "Polysaccharide lyases," *Appl. Biochem. Biotech.* 12(2): 135-176 (1986).

LINN et. al., "Isolation and characterization of two chondroitin lyases from *Bacteroides thetaiotaomicron*," *J. Bacteriol.* 156(2):859-866 (1983).

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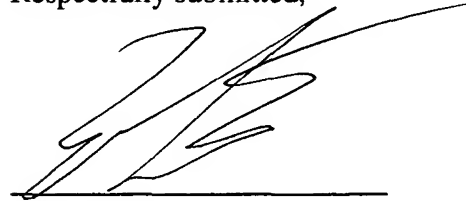
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ZAWADZKI, et al., "Blockade of metastasis formation by CD44-receptor globulin," *Int. J. Cancer* 75(6):919-924 (1998).

**Remarks**

This statement should not be interpreted as a representation that an exhaustive search has been conducted or that no better art exists. Moreover, Applicants invite the Examiner to make an independent evaluation of the cited art to determine its relevance to the subject matter of the present application. Applicants are of the opinion that their claims patentably distinguish over the art referred to herein, either alone or in combination.

Respectfully submitted,

A handwritten signature in black ink, appearing to be 'Zhaoyang Li', written over a horizontal line.

Zhaoyang Li  
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Dated: July 18, 2003

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						Filing Date	July 18, 2003
						First Named Inventor	Elizabeth M. Denholm
						Group Art Unit	
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## U.S. PATENT DOCUMENTS

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## FOREIGN PATENT DOCUMENTS

Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T°
		Office. <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
		JP	51075042		Ono Pharm Co. Ltd.	06-29-1979		
		PCT	WO 96/01894		IBEX Technologies	01-25-1996		
		PCT	WO 96/01648		IBEX Technologies	01-25-1996		
		PCT	WO 96/08559		Cardiac CRC Nominees PTY, Ltd.	03-21-1996		

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OTHER ART -- NON PATENT LITERATURE DOCUMENTS			
Examiner's Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
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